## **WE CLAIM**:

- 1. A use of a propargylamine to enhance the activity of an antineoplastic drug.
- 2. A use according to claim 1 wherein the propargylamine increases the sensitivity of a tumor to the antineoplastic drug.
  - 3. A use according to claim 2 wherein the tumor is a drug resistant tumor.
  - 4. A use according to claim 1 wherein the propargylamine protects normal cells from the cytotoxic effects of the antineoplastic drug.
- 10 5. A use of (a) a propargylamine and (b) an antineoplastic drug to treat cancer.
  - 6. A use according to any one of claims 1-5 wherein the propargylamine is of the general formula I

15  $R''(CR'_2)_xC - N$   $R_2$   $R_2$   $R_3$   $R_2$   $R_3$   $R_3$ 

20 x is an integer ranging from 0 to 13; y is an integer ranging from 0 to 5;

z is an integer ranging from 0 to 5;

 $R_1$ ,  $R_2$  and  $R_3$  are the same or different and represent hydrogen or a straight chain or branched lower alkyl; and

25 R' and R' are the same or different and represent hydrogen, phenyl or a halogen and pharmaceutically acceptable salts thereof.

7. A use according to claim 6 wherein y is 1.

8. A use according to claim 7 wherein the propargylamine is R-2-heptyl-methyl propargylamine (R-2HMP).

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- 9. A use according to claim 7 wherein the propargylamine is selected from the group consisting o f N-(1-Propyl) N-methylpropargylamine; N-(2-Propyl) N-methylpropargylamine; N-methylpropargylamine; N-(1-Butyl)N-(1-Pentyl) N-methylpropargylamine; N-(1-Hexyl) N-methylpropargylamine; N-(1-Heptyl) N-methylpropargylamine; N-(1-Octyl)N-methylpropargylamine; N-(1-Nonyl) N-methylpropargylamine; N-(1-Decyl) N-methylpropargylamine; N-(1-Undecyl) N-methylpropargylamine; N-(1-Dodecyl) N-methylpropargylamine; (R)-N-(2-Butyl)N-methylpropargylamine;(R)-N-(2-Pentyl) N-methylpropargylamine;(R)-N-(2-Hexyl) N-methylpropargylamine; 15 (R)-N-(2-Heptyl)N-m/ethylpropargylam in e; (R)-N-(2-Octyl) N-methylpropargylamine; (R)-N-(2-Octyl) N-methylpropargylamine; (R)-N-(2-Decyl)N-methylpropargylamine; (R)-N-(2-Un decyl)N-methylpropa/tgylamine; an,d (R)-N-(2-Dodecyl)
- 20 10. A use according to claim 6 wherein y is 0.

N-methylpropargylamine.

- 11. A use according to claim 10 wherein the propargylamine is R-2-heptyl-propargylamine R-2HPA).
- 12. A use according to claim 10 wherein said propargylamine is selected from the group consisting of N-(1-Propyl) propargylamine; N-(2-Propyl) propargylamine; N-(1-Butyl) propargylamine; N-(1-Pentyl) propargylamine; N-(1-Hexyl) propargylamine; N-(1-Heptyl) propargylamine; N-(1-Nonyl)

propargylamine; N-(1-Decyl) propargylamine; N-(1-Undecyl) propargylamine; N-(1-Dodecyl) propargylamine; (R)-N-(2-Pentyl) propargylamine; (R)-N-(2-Heptyl) propargylamine; (R)-N-(2-Octyl) propargylamine; (R)-N-(2-Octyl) propargylamine; (R)-N-(2-Decyl) propargylamine; (R)-N-(2-Undecyl) propargylamine; and (R)-N-(2-Dodecyl) propargylamine.

- 13. A use according to any one of claims 1 to 7, 9, 10 or 12 wherein the propargylamine is a chiral compound and is the R
  10 enantiomer.
  - 14. A use according to any one of claims 1-6 wherein the propargylamine is R-deprenyl.
  - 15. A use according to any one of claims 1-6 wherein the propargylamine is R-desmethyldeprenyl.
- 15 16. A use according to any one of claims 1-5 wherein the propargylamine is Rasagiline.
  - 17. A use according to any one of claims 1-16 wherein the animal is a human.
- 18. A use according to any one of claims 1-17 wherein the 20 antineoplastic drug is selected from the group consisting of cytosine arabinoside, cis-platinum, cyclophospamide, adriamycin, daunomycin, and 5-fluorouracil.
- 19. A pharmaceutical composition for enhancing the activity of an antineoplastic drug comprising an effective amount of a propargylamine in admixture with a suitable diluent or carrier.

- 20. A pharmaceutical composition according to claim 19 for increasing the sensitivity of a tumor to the antineoplastic drug.
- 21. A pharmaceutical composition according to claim 19 for protecting normal cells from the cytotoxic effects of the antineoplastic drug.
- 22. A pharmaceutical composition for treating cancer comprising an antineoplastic drug and an effective amount of a propargylamine.
- 23. A pharmaceutical composition according to any one of claims 19 to 22, wherein the propargylamine is of the general formula I:

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$$R_{1} \quad (CH_{2})_{y}H$$

$$R''(CR'_{2})_{x}C \longrightarrow N$$

$$R_{2} \quad (CH_{2})_{z}C \equiv CR_{3}$$

15 wherein

x is an integer ranging from 0 to 13; y is an integer ranging from 0 to 5; z is an integer ranging from 0 to 5;

 $R_1$ ,  $R_2$  and  $R_3$  are the same or different and represent

20 hydrogen or a straight chair or branched lower alkyl; and

R' and R' are the same or different and represent hydrogen, phenyl or a halogen and pharmaceutically acceptable salts thereof.

- 24. A pharmaceutical composition according to claim 23 wherein y is 1.
- 25 25. A pharmaceutical composition according to claim 24 wherein the propargylamine is R-2-heptyl-methyl propargylamine (R-2HMP).

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- A pharmaceutical composition according to claim 24 wherein 26. the propargylamine is selected from the group consisting of N-(1-Propyl) N-methylpropargylamine; N-(2-Propyl) N-methylpropargylamine; N-(1-Butyl)N-methylprdpargylamine; N-(1-Pentyl) N-methylpropargylamine; N-(1 Hexyl) N-methylpropargylamine; N-(1-Heptyl) N-methylpropargylamine; N-(1-Octyl)N-methylpropargylamine; N-(1-Nonyl) N-methylpropargylamine; N-(1-Decyl) N-methylpropargylamine; N-(1-Un decvl) N-methylpropargylamine; N-(1-podecyl) N-methylpropargylamine; (R)-N-(2-Butyl)N-methylpropargylamine; (R)-N-(2-Pentyl)N-methylpropargylamine; (R)-N-(2-Hexyl) N-methylpropargylamine; (R)-N-(2-Heptyl)N-methylpropargylamine; (R)-N-(2-Octv1)N-methylpropargylamine; (R)-N-(2-Octyl) N-methylpropargylamine; (R)-N-(2-Decvl) N-methylpropargylamine; (R)-N-(2-Un decyl)N-methylpropargylamine; and (R)-N-(2-Dodecyl) N-methylpropargylamine.
- 27. A pharmaceutical composition according to claim 23, wherein y is 0.
- 28. A pharmaceutical composition according to claim 27 wherein 20 the propargylamine is R-2-heptyl-propargylamine (R-2HPA).
- 29. A pharmaceutical composition according to claim 27 wherein said propargylamine is selected from the group consisting of N-(1-Propyl) N-(2-Propyl) propargylamine; propargylamine; N-(1-Butyl) propargylamine; N-(1-Fentyl) propargylamine; N-(1-Hexyl) 25 propargylamine; N-(1-Heptyl) propargylamine; N-(1-Octyl) N-(1-|Nonyl) propargylamine; propargylamine; N-(1-Decyl) propargylamine; N-(1-Undecyl) propargylamine; N-(1-Dodecyl) propargylamine; (R)-N-(2-Pentyl) propargylamine; (R)-N-(2-Pentyl) propargylamine; (R)-N-(2-Hexyl) propargylamine; (R)-N-(2-Heptyl)

propargylamine; (R)-N-(2-Octyl) propargylamine; (R)-N-(2-Octyl) propargylamine; (R)-N-(2-Decyl) propargylamine; (R)-N-(2-Undecyl) propargylamine; and (R)-N-(2-Dødecyl) propargylamine.

- 30. A pharmaceutical composition according to any one of claims
  19 to 24, 26, 27 or 29 wherein the propargylamine is a chiral compound and is the R-enantiomer.
  - 31. A pharmaceutical composition according to any one of claims 19 to 23, wherein the propargy amine is R-deprenyl.
- 32. A pharmaceutical composition according to any one of claims 10 19 to 23, wherein the propargylamine is R-desmethyldeprenyl.
  - 33. A pharmaceutical composition according to any one of claims 19 to 22, wherein the propargylamine is Rasagiline.
  - 34. A method for enhancing the activity of an antineoplastic drug comprising administering an effective amount of a propargylamine to an animal in need thereof.
  - 35. A method according to claim 34 wherein the propargylamine increases the sensitivity of a tumor to an antineoplastic drug.
  - 36. A method according to claim 35 wherein the tumor is a drug resistant tumor.
- 20 37. A method according to claim 34 wherein the propargylamine protects normal cells form the cytotoxic effects of the antineoplastic drug.

- 38. A method for treating cancer comprising administering an antineoplastic drug and an effective amount of a propargylamine to an animal in need thereof.
- 39. A method according to any one of claims 34 to 38, wherein the propargylamine is of the general formula I

$$R_{1} \quad (CH_{2})_{y}H$$

$$| \quad /$$

$$R''(CR'_{2})_{x}C - N$$

$$| \quad \backslash$$

$$R_{2} \quad (CH_{2})_{z}C \equiv CR$$

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wherein

x is an integer ranging from 0 to 13;

y is an integer ranging from 0 to 5;

z is an integer ranging from 0 to 5;

15 R<sub>1</sub>, R<sub>2</sub> and R<sub>3</sub> are the same or different and represent hydrogen or a straight chain or branched lower alkyl; and

R' and R" are the same or different and represent hydrogen, phenyl or a halogen and pharmaceutically acceptable salts thereof.

40. A method according to claim 39 wherein y is 1.

- 20 41. A method according to claim 40 wherein the propargylamine is R-2-heptyl-methyl propargylamine (R-2HMP).
  - 42. A method according to claim 39 wherein the propargylamine is selected from the group consisting of N-(1-Propyl) N-methylpropargylamine; N-(2-Propyl) N-methylpropargylamine; N-(1-Butyl) N-methylpropargylamine; N-(1-Pentyl) N-methylpropargylamine; N-(1-Hexyl) N-methylpropargylamine; N-(1-Hexyl) N-methylpropargylamine;

N-methylpropargylamine; N-(1-Hexyl) N-methylpropargylamine; N-(1-Heptyl) N-methylpropargylamine; N-(1-Octyl)

N-methylpropargylamine; N-(1+Nonyl) N-methylpropargylamine; N-(1-Decyl) N-methylpropargylamine; N-(1-Undecvl) N-methylpropargylamine; N-(1-Dodecyl) N-methylpropargylamine; (R)-N-(2-Butyl)N-methylpropargylamine; (R)-N-(2-Pentyl)N-methylpropargylamine; (R)-N-(2-Hexyl) N-methylpropargylamine; (R)-N-(2-Heptyl)N-methylpropargylamine; (R)-N-(2-Octyl)N-methylpropargylamine; (R)-N-(2-Octyl) N-methylpropargylamine; (R)-N-(2-Decyl)N-methylpropargylamine; (R)-N-(2-Un decv1)N-methylpropargylamine; and (R)-N-(2-Dodecvl)N-methylpropargylamine. 10

- 43. A method according to claim 39, wherein y is 0.
- 44. A method according to claim 43 wherein the propargylamine is R-2-heptyl-propargylamine (R-2 HPA).
- 45. A method according to claim 43 wherein the propargylamine 15 is selected from the group consisting of N-(1/Propyl) propargylamine; N-(2-Propyl) propargylamine; N-(1-Butyl) propargylamine; N-(1-Pentyl) propargylamine; N-(1-Hexy)propargylamine; N-(1-Heptyl) propargylamine; N-(1-Octyl) propargylamine; N-(1-Nonyl) propargylamine; N-(1-Decyl) propargylamine; N-(1-Undecyl) 20 propargylamine; N-(1-Dodecyl) propargylamine; (R)-N-(2-Butyl) propargylamine; (R)-N-(2-Pentyl) propargylamine; (R)-N-(2-Hexyl) propargylamine; (R)-N-(2-Heptyl) propargylamine; (R)-N-(2-Octyl) propargylamine; (R)-N-(2-Octyl) propargylamine; (R)-N-(2-Decyl) propargylamine; (R)-N-(2-Undecyl) propargylamine; and (R)-N-(2-Dodecyl) 25 propargylamine.
  - 46. A method according to any one of claims 34 to 39, wherein the propargylamine is R-deprenyl.

- 47. A method according to any one of claims 34 to 39, wherein the propargylamine is R-desmethyldeprenyl.
- 48. A method according to any one of claims 34 to 38, wherein the propargylamine is Rasagiline.
- 5 49. A method according to any one of claims 34 to 48, wherein the animal is a human.
  - A method according to any one of claims 34 to 49 wherein the antineoplastic drug is selected from the group consisting of cytosine arabinoside, cis-platinum, cyclophospamide, adriamycin, daunomycin, and 5-fluorouracil.
  - 51. A method according to any one of claims 34 to 40, 42, 43 and 45 wherein the propargylamine is a chiral compound and is the Renantiomer.